In re Application of: Khan, Saeed R. PATENT
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Amendments to the Claims:

Please amend claims 2-9 and 11-19 as shown in the listing of the claims.

Please cancel claim 1 without prejudice.

This listing of the claims will replace all prior listing of the claims in this application.

#### Listing of the Claims:

- 1. (Canceled).
- 2. (Currently Amended) The compound of claim [4] 19, wherein

 $L^1$  is a bond, substituted or unsubstituted  $C_1$ - $C_{20}$  alkylene, or substituted or unsubstituted 2 to 20 membered heteroalkylene;

 $L^2$  is independently a bond, substituted or unsubstituted  $C_1$ - $C_{20}$  alkylene, substituted or unsubstituted 2 to 20 membered heteroalkylene, or substituted or unsubstituted 3 to 8 membered heteroevoloalkylene; and

 $R^2$ ,  $R^3$ , and  $R^4$  are each independently hydrogen, substituted or unsubstituted  $C_1$ - $C_{20}$  alkyl, substituted or unsubstituted 2 to 20 membered heteroalkyl, substituted or unsubstituted  $C_3$ - $C_8$  cycloalkyl, substituted or unsubstituted 3 to 8 membered heterocycloalkyl, substituted or unsubstituted or unsubstituted heteroaryl.

- (Currently Amended) The compound of claim [1] 19, wherein
  - $\mathrm{L}^1$  is a bond, unsubstituted  $\mathrm{C}_1\text{-}\mathrm{C}_{10}$  alkylene, or unsubstituted 2 to 10 membered heteroalkylene;
  - $L^2$  is independently a bond, unsubstituted  $C_1$ - $C_{10}$  alkylene, unsubstituted 2 to 10 membered heteroalkylene, or unsubstituted 3 to 8 membered heteroeyeloalkylene; and

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R<sup>2</sup> R<sup>3</sup>, and R<sup>4</sup> are each independently hydrogen, substituted or unsubstituted C<sub>1</sub>-C<sub>10</sub> alkyl, substituted or unsubstituted 2 to 10 membered heteroalkyl, substituted or unsubstituted C<sub>3</sub>-C<sub>8</sub> cycloalkyl, substituted or unsubstituted 3 to 8 membered heterocycloalkyl. substituted or unsubstituted arvl, or substituted or unsubstituted heteroarvl.

4 (Currently Amended) The compound of claim [4] 19, wherein

> L1 is a bond, substituted or unsubstituted C1-C10 alkylene, or substituted or unsubstituted 2 to 10 membered heteroalkylenef+1

L2 is independently a bond, substituted or unsubstituted C1-C10 alkylene, substituted or unsubstituted 2 to 10 membered heteroalkylene, or substituted or unsubstituted 3 to 8 membered heterocycloalkylene.

5. (Currently Amended) The compound of claim [4] 19, wherein R1 has the formula:

wherein:

L2 is a bond, unsubstituted C1-C10 alkylene, unsubstituted 2 to 10 membered heteroalkylene, or unsubstituted 3 to 8 membered heterocycloalkylene:

R<sup>2</sup> is R<sup>21</sup>-substituted or unsubstituted C<sub>1</sub>-C<sub>15</sub> alkyl, R<sup>21</sup>-substituted or unsubstituted 2 to 10 membered heteroalkyl, R21-substituted or unsubstituted C3-C8 cycloalkyl, R21-substituted or unsubstituted 3 to 8 membered heterocycloalkyl, R21-substituted or unsubstituted aryl, or R21-substituted or unsubstituted heteroaryl, wherein:

R<sup>21</sup> is a halogen. -OH, -SH, -NH<sub>2</sub>, -CF<sub>3</sub>, -B(OH)<sub>2</sub>, -C(0)NHOH, unsubstituted C<sub>1</sub>-C<sub>10</sub> alkyl, unsubstituted 2 to 10 membered heteroalkyl, unsubstituted C3-C8 cycloalkyl.

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unsubstituted 3 to 8 membered heterocycloalkyl, unsubstituted aryl, unsubstituted heteroaryl, or -OR<sup>22</sup>, wherein:

 $R^{22}$  is unsubstituted  $C_3$ - $C_{10}$  alkyl, unsubstituted 2 to 10 membered heteroalkyl, unsubstituted  $C_3$ - $C_8$  cycloalkyl, unsubstituted 3 to 8 membered heterocycloalkyl, unsubstituted aryl, unsubstituted aryl, unsubstituted heteroaryl, or -(CH<sub>2</sub>)<sub>q</sub>B(OH)<sub>2</sub>, wherein q is an integer from 1 to 5.

6. (Currently Amended) The compound of claim [1] 19, wherein R<sup>1</sup> has the formula:

wherein:

# L2 is a bond, or unsubstituted 3 to 8 membered heterocycloalkylene;

 $R^2$  is  $R^{21}$ -substituted or unsubstituted  $C_1$ - $C_{15}$  alkyl, or  $R^{21}$ -substituted or unsubstituted aryl, wherein:

R<sup>21</sup> is halogen, -OH, -SH, -NH<sub>2</sub>, -CF<sub>3</sub>, -B(OH)<sub>2</sub>, -C(0)NHOH, unsubstituted C<sub>1</sub>-C<sub>10</sub> alkyl, unsubstituted 2 to 10 membered heteroalkyl, unsubstituted C<sub>3</sub>-C<sub>8</sub> cycloalkyl, unsubstituted 3 to 8 membered heterocycloalkyl, unsubstituted aryl, unsubstituted heteroaryl, or -OR<sup>22</sup>, wherein;

 $R^{22}$  is unsubstituted  $C_1$ - $C_{10}$  alkyl, or unsubstituted 2 to 10 membered heteroalkyl.

7. (Currently Amended) The compound of claim 6, wherein

## L2 is a bond or piperazinylene;

R<sup>2</sup> is unsubstituted C<sub>1</sub>-C<sub>15</sub> alkyl, or R<sup>21</sup>-substituted or unsubstituted aryl, wherein:

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 $R^{21}$  is halogen, -C(0)NHOH, unsubstituted  $C_1$ - $C_{10}$  alkyl, unsubstituted 2 to 10 membered heteroalkyl, or -OR<sup>22</sup>, wherein:

R<sup>22</sup> is unsubstituted C<sub>1</sub>-C<sub>10</sub> alkyl, or unsubstituted 2 to 10 membered heteroalkyl.

8. (Currently Amended) The compound of claim [4] 19, wherein R<sup>1</sup> has the formula:

wherein:

 $R^4$  is unsubstituted  $C_1$ - $C_10$  alkyl, unsubstituted 2 to 10 membered heteroalkyl, unsubstituted  $C_3$ - $C_8$  cycloalkyl, unsubstituted 3 to 8 membered heterocycloalkyl, unsubstituted aryl, or unsubstituted heteroaryl;

 $R^3$  is  $R^{31}$ -substituted or unsubstituted  $C_{1}$ - $C_{10}$  alkyl,  $R^{31}$ -substituted or unsubstituted 2 to 10 membered heteroalkyl, R -substituted or unsubstituted  $C_{3}$ - $C_{8}$  cycloalkyl,  $R^{31}$ -substituted or unsubstituted 3 to 8 membered heterocycloalkyl,  $R^{31}$ -substituted or unsubstituted aryl, or  $R^{31}$ -substituted or unsubstituted heteroaryl, wherein:

R<sup>31</sup> is halogen, -OH, -SH, -NH<sub>2</sub>, -CF<sub>3</sub>, -B(OH)<sub>2</sub>, -C(0)NHOH, unsubstituted C<sub>1</sub>-C<sub>10</sub> alkyl, unsubstituted 2 to 10 membered heteroalkyl, unsubstituted C<sub>3</sub>-C<sub>8</sub> cycloalkyl, unsubstituted 3 to 8 membered heterocycloalkyl, unsubstituted aryl, unsubstituted heteroaryl, or -OR<sup>32</sup>, wherein:

R<sup>32</sup> is unsubstituted C<sub>1</sub>-C<sub>10</sub> alkyl, unsubstituted 2 to 10 membered heteroalkyl, unsubstituted C<sub>3</sub>-C<sub>8</sub> cycloalkyl, unsubstituted 3 to 8 membered heterocycloalkyl,

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unsubstituted aryl, unsubstituted heteroaryl, or -(CH2)<sub>m</sub>B(OH)<sub>2</sub>, wherein m is an integer from 1 to 5.

9. (Currently Amended) The compound of claim [1] 19, wherein R1 has the formula:

wherein:

R<sup>4</sup> is unsubstituted C<sub>1</sub>-C<sub>10</sub> alkyl;

R3 is R31-substituted aryl, wherein:

 $R^{31}$  is a halogen, -C(0)NHOH, unsubstituted  $C_1$ - $C_{10}$  alkyl, unsubstituted 2 to 10 membered heteroalkyl, or -O $R^{32}$ , wherein:

 $R^{32}$  is an unsubstituted  $C_1$ - $C_{10}$  alkyl, unsubstituted 2 to 10 membered heteroalkyl, or -(CH<sub>2</sub>)<sub>m</sub>B(OH)<sub>2</sub>, wherein m is 1 to 5.

- 10. (Canceled).
- 11. (Currently Amended) The compound of claim [4] 19, wherein R<sup>1</sup> is para to -L<sup>1</sup>-B(OH)<sub>2</sub>.
- 12. (Currently Amended) The compound of claim [4] 19, wherein X is -C=.
- 13. (Currently Amended) The compound of claim [4] 19, wherein L<sup>1</sup> is a bond or methylene.

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14. (Withdrawn/Currently Amended) A method of treating a tumor or cancer in a patient in need thereof, the method comprising the step of administering to said patient an effective amount of a compound having the Formula:

wherein

X1 is O or N=:

 $\mathbf{L}^{1}$  is a bond, substituted or unsubstituted alkylene, or substituted or unsubstituted heteroalkylene; and

R1 has the formula:

$$(1) - \frac{\frac{5}{5} - L^{2} - C - L^{3} - R^{2}}{5}$$

$$(2) - \frac{\frac{5}{5} - L^{4} - \frac{1}{5} - R^{3}}{\frac{5}{5} - L^{4} - \frac{1}{5} - R^{3}}$$

$$(2) - \frac{\frac{5}{5} - L^{4} - \frac{1}{5} - R^{3}}{\frac{5}{5} - L^{4} - \frac{1}{5} - R^{3}}$$

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#### wherein

n is 0 or 1.

X2 is N(R4) or CH(R4):

L2.L3.L4, and L5 are independently a bond, substituted or unsubstituted alkylene, substituted or unsubstituted heteroalkylene, or substituted or unsubstituted heterocycloalkylene: and

R2-R3- and R4- are independently hydrogen, substituted or unsubstituted alkyl, substituted or unsubstituted heteroalkyl, substituted or unsubstituted cycloalkyl, substituted or unsubstituted heterocycloalkyl, substituted or unsubstituted arvl, or substituted or unsubstituted heteroaryl.

wherein if X is -C=, L2 is a bond, L3 is unsubstituted alkylene, R1 is (1), and R4 is para to-L4-B(OH): then R2 is not substituted or unsubstituted anyl or substituted or unsubstituted heteroarvi of claim 19.

- 15 (Withdrawn/Currently Amended) The method of claim 14, wherein said tumor is selected from the group consisting of breast, cervical, stomach, colon, bladder, rectal, liver, pancreatic, lung, cervix uteri, corpus uteri, ovary, prostate, testis, renal, brain/cns. head, neck, throat, anal and oral cancers, eye or ocular cancer, skin melanoma. Ewing's Sarcoma, Kaposi's Sarcoma, basal cell carcinoma and squamous cell carcinoma, small cell lung cancer, mouth/pharynx, esophageal, larynx, kidney and lymphoma, acute lymphocytic leukemia, and acute myelogenous leukemia.
- 16 (Withdrawn/Currently Amended) A method of inhibiting MDM2 expression in a mammal, the method comprising the step of administering an amount of a compound of claim 19 effective to inhibit said expression, said compound having the Formula:

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### wherein

X1-is-O or-N=;

 $\mathbf{L}^{1}$  is a bond, substituted or unsubstituted alkylene, or substituted or unsubstituted heteroalkylene; and

## R+has the formula:

$$(1) \qquad \begin{array}{c} \underbrace{\frac{1}{5} - 1^{2} - 1^{3} - R^{2}}_{5} \\ \underbrace{\frac{1}{5} - 1^{4} - 1^{5} - R^{3}}_{7} \\ \underbrace{\frac{1}{5} - 1^{4} - 1^{5} - R^{3}}_{7} \\ \underbrace{\frac{1}{5} - 1^{4} - 1^{5} - R^{3}}_{7} \\ \end{array}$$

### wherein

n is 0 or 1;

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L<sup>2</sup>, L<sup>4</sup>, rand L<sup>5</sup> are independently a bond, substituted or unsubstituted alkylene, substituted or unsubstituted heteroalkylene, or substituted or unsubstituted heteroeycloalkylene; and

 $R^2$ ,  $R^3$ , and  $R^4$  are independently hydrogen, substituted or unsubstituted alkyl, substituted or unsubstituted heteroalkyl, substituted or unsubstituted exploalkyl, substituted or unsubstituted aryl, or substituted or unsubstituted aryl, or substituted or unsubstituted heteroaryl.

wherein if X is -C ,  $L^2$  is a bond,  $L^2$  is unsubstituted alkylene,  $R^1$  is (1), and  $R^2$  is para to  $-L^1$ -B(OH)<sub>2</sub>, then  $R^2$  is not substituted or unsubstituted aryl or substituted or unsubstituted heteroaryl.

17. (Currently Amended) The compound of claim [4] 19, wherein R<sup>1</sup> has the formula:

$$(2) \qquad \qquad (3) \qquad (3)$$

 (Currently Amended) The compound of claim [‡] 19, wherein the compound having the formula:

has formulae:

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$$(HO)_2B$$

wherein R' is halogen, alkyl, heteroalkyl, alkoxy, heteroalkoxy, or hydroxamic acid.

#### 19. (Currently Amended) A compound having formula:

wherein

 $X^1$  is C= or -N=;

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 $\mathbf{L}^1$  is a bond, substituted or unsubstituted alkylene, or substituted or unsubstituted heteroalkylene;

## R1 has the formula:

wherein

(2)

n is 0 or 1;

 $X^2$  is  $-N(R^4)$  -N- or  $-CH(R^4)$  -CH-: and

 $R^2$ ,  $R^3$ , and  $R^4$  are <u>each</u> independently hydrogen, substituted or unsubstituted alkyl, substituted or unsubstituted heteroalkyl, substituted or unsubstituted cycloalkyl, substituted or unsubstituted aryl, or substituted or unsubstituted heteroaryl.